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Declaration under Rule 4.17:

as to applicant's entitlement to apply for and be granted a patent (Rule 4.17(ii)) for the following designations AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ. CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM. PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, ARIPO patent (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT. LU, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG)

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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.



(57) Abstract: According to the invention there is provided the use of a low molecular weight thrombin inhibitor, or a pharmaceutically acceptable derivative thereof, for the manufacture of a medicament for use in cholesterol-lowering therapy and/or modification of lipid (triglyceride), lipoprotein, and apolipoprotein, profiles associated with an increased risk of cardiovascular complications.



Intermonal application No. PCT/SE 2004/000417

A. CLASSIFICATION OF SUBJECT MATTER IPC7: A61K 38/05 A61D 7/02 A61D 3/0

IPC7: A61K 38/05, A61P 7/02, A61P 3/06, A61P 3/10
According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC7: A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

SE,DK,FI,NO classes as above

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

CHEM. ABS DATA, WPI DATA, EMBASE, MEDLINE, BIOSIS

C. DOCUMENTS CONSIDERED TO BE RELEVANT

| Category* | Citation of document, with indication, where appropriate, of the relevant passages | Relevant to claim No. | |
|-----------|--|-----------------------|--|
| x | EP 0432537 A2 (IKETON FARMACEUTICI S.R.L.), 19 June 1991 (19.06.1991), see page 2, lines 29-32 and page 4, lines 25-26 | 1-26,42-44 | |
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| Х | WO 9811896 A1 (MERCK & CO., INC.), 26 March 1998 (26.03.1998), see the claims and page 22, 1 ines 27-29 | 27-41 | |
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| | Further documents are listed in the continuation of Box | K.C. X See patent family annex. | | |
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| "P" | Special categories of cited documents: document defining the general state of the art which is not considered to be of particular relevance earlier application or patent but published on or after the international filling date document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) document referring to an oral disclosure, use, exhibition or other means document published prior to the international filing date but later than the priority date claimed | "X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is considered to involve an inventive step when the document is combined with once or more other such documents such combinates. | | |
| | of the actual completion of the international search | Date of mailing of the international search report | | |
| | June 2004 11 3 -09- 2004 | l 1 6 − 09− 2004 | | |
| Swe Box Facs | te and mailing address of the ISA/ dish Patent Office 5055, S-102 42 STOCKHOLM dimile No. +46 8 666 02 86 PCT/ISA/210 (second sheet) | Authorized officer Carolina Gómez Lagerlöf/EÖ Telephone No. + 46 8 782 25 00 | | |

International application No.
PCT/SE 2004/000417

| This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons: 1. | Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet) | | | | | | |
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| because they relate to subject matter not required to be searched by this Authority, namely: see next sheet* 2. Claims Nos.: 1-2,27 because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically: see next sheet* 3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a). Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet) This International Searching Authority found multiple inventions in this international application, as follows: 1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims. 2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee. 3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.: | This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons: | | | | | | |
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| Remark on Protest The additional search fees were accompanied by the applicant's protest. | | | | | | | |
| No protest accompanied the payment of additional search fees. | No protest accompanied the payment of additional search fees. | | | | | | |

International application No.
PCT/SE 2004/000417

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Claims15-26, 43 relate to methods of treatment of the human or animal body by surgery or by therapy or diagnostic methods practised on the human or animal body (PCT Rule 39.1(iv)). Nevertheless, a search has been executed for these claims. The search has been based on the alleged effects of the compounds or compositions.

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Present claims 1-2 and 27 relate to compounds defined by reference to a desirable characteristic or property, namely low molecular weight thrombin inhibitor, cholesterol-lowering or lipid-lowering therapeutic agents. The claims cover the use of all compounds having this characteristic or property, whereas the application provides support within the meaning of Article 6 PCT or disclosure within the meaning of Article 5 PCT for only a very limited number of such. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Independent of the above reasoning, the claims also lack clarity (Article 6 PCT). An attempt is made to define the compound by reference to a result to be achieved. Again, this lack of clarity in the present case is such as to render a meaningful search over the whole of the claimed scope impossible.

Consequently, the search has been carried out for those parts of the claims which appear to be clear, supported and disclosed, namely those parts relating to compounds defined in claims 4-13. A search for the general wording "low molecular weight thrombin inhibitor" have also been made.

Inter al application No. PCT/SE 2004/000417

| EP | 0432537 | A2 | 19/06/1991 | SE AT DE ES IE IT IT JP PT US | 0432537 T3 117697 T 69016383 D,T 2068969 T 904225 A 1237518 B 8922504 D 3210302 A 95961 A,B 5164378 A | 15/02/1995 13/07/1995 01/05/1995 05/06/1991 08/06/1993 00/00/0000 13/09/1991 13/09/1991 17/11/1992 |
|----|---------|----|------------|--|--|--|
| WO | 9811896 | A1 | 26/03/1998 | AU CA EP GB JP US US | 723315 B 4350897 A 2265827 A 0946178 A 9621970 D 2001500875 T 6235706 B 6251852 B 6403571 B 2001036913 A | 24/08/2000 14/04/1998 26/03/1998 06/10/1999 00/00/0000 23/01/2001 22/05/2001 26/06/2001 11/06/2002 01/11/2001 |